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FLORIDA STATE UNIV TALLAHASSEE DEPT OF STATISTICS

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TESTING WHETHER ONE REGRESSION FUNCTION IS EVERYWHERE LARGER TH--ETC(U)

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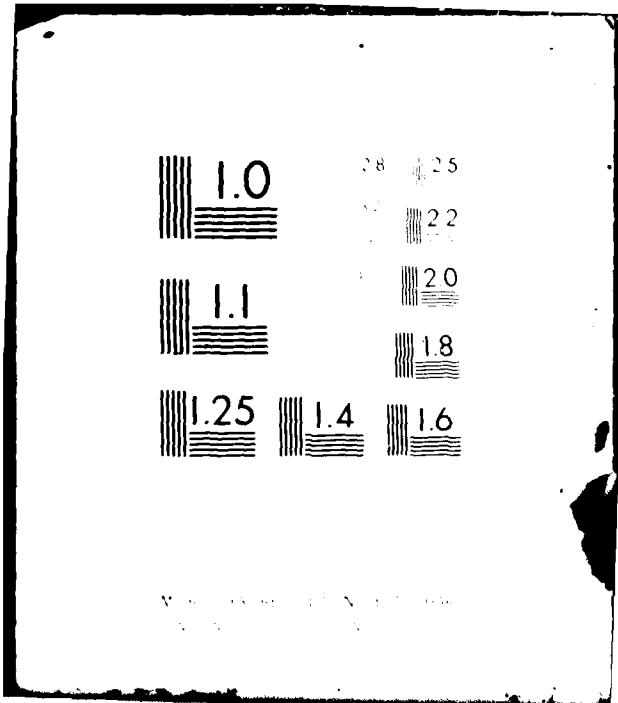
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TESTING WHETHER ONE REGRESSION  
FUNCTION IS EVERYWHERE LARGER THAN ANOTHER

by

Roger L. Berger

FSU Statistics Report M607  
USARO Technical Report D- 52

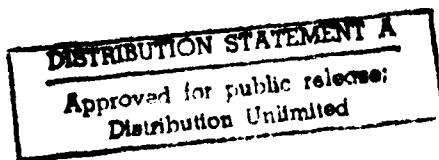
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February, 1982



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This research was supported in part by the U.S. Army Research Office  
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## SUMMARY

The problem of testing whether one regression function is larger than another on a specified compact set  $R$  is considered. The regression functions must be linear functions of the parameters but need not be linear functions of the independent variables. The proposed test statistic is compared to a standard  $t$  percentile. The test has an exactly specified size in typical situations. Properties of the power function of the test are investigated. The related question of comparing a regression function to a specified function is also considered.

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### 1. Introduction

In this paper we consider testing whether the regression function from one population is everywhere above the regression function from another population. A medical researcher, for example, might be interested in testing whether the mean of a response variable in a diseased population is larger than the mean of a response variable in a healthy population for all possible values of an independent variable. Examples such as these have been discussed by Tsutakawa and Hewett (1978), Hewett and Lababidi (1980) and Spurrier, Hewett and Lababidi (1980) who have considered this testing problem.

The model considered herein generalizes the models of Tsutakawa and Hewett (1978) and Hewett and Lababidi (1980) in two ways. First the regression functions may be functions other than linear functions of the independent variables. For example, the regression functions may be quadratic or higher degree polynomials. Second, the two regression functions need not be of the same functional form. For example, one may be assumed to be a linear function and the other a quadratic function. The test proposed herein reduces to the tests of Tsutakawa and Hewett (1978) and Hewett and Lababidi (1980) for the special models they consider. The proposed test requires no new tables for its implementation. Only a standard  $t$  table is needed.

The model and test are presented in Section 2.1. A numerical example using the organ weight data of normal and diabetic mice is presented in Section 2.2. Properties of the test, including its power, are discussed in Section 3. The related problem of testing whether a single regression function everywhere exceeds a specified function is discussed in Section 4.

## 2. Model, Test and Application

### 2.1. Model and Test

Let  $\{(X_{1j}, Y_{1j}), j = 1, \dots, n_1\}$  and  $\{(X_{2j}, Y_{2j}), j = 1, \dots, n_2\}$  denote two independent sets of observations where

$X_{ij} = (X_{ij1}, \dots, X_{ijk})$ . The  $X_{ij}$  may be observed random vectors or design variables fixed by the experimenter. The entire analysis is conditioned on the observed values of  $X_{ij}$ . Let  $R$  denote a closed and bounded subset of  $k$ -dimensional Euclidean space.  $R$  is the set of possible values of the independent variables  $X_{ij}$ . We assume that given the  $X_{ij}$  the  $Y_{ij}$  are independent normal random variables with

$$E(Y_{ij} | X_{ij1} = x_{ij1}, \dots, X_{ijk} = x_{ijk}) = \sum_{m=1}^{p_i} \beta_{im} f_{im}(x_{ij1}, \dots, x_{ijk}) = f_i(x_{ij}) \beta_i$$

and

$$\text{var}(Y_{ij} | X_{ij1} = x_{ij1}, \dots, X_{ijk} = x_{ijk}) = \sigma^2.$$

$\beta_i = (\beta_{i1}, \dots, \beta_{ip_i})'$ ,  $i = 1, 2$ , and  $\sigma^2$  are unknown parameters.

The  $f_i(x) = (f_{i1}(x), \dots, f_{ip_i}(x))$ ,  $i = 1, 2$ , are known vectors of functions which define the functional form of the regression functions. For example the  $f_{ij}$  might be polynomials such as  $1, x_1, x_2^2$ , or  $x_1x_3$ . By allowing  $p_1 \neq p_2$  and  $f_{1m} \neq f_{2m}(x)$ , this model allows the two regression functions to have different functional forms. The first might be a linear function and the second a quadratic function. But usually  $p_1 = p_2$  and  $f_1(x) = f_2(x)$  will be chosen so the regression functions have the same functional form.

We wish to compare the regression functions  $f_1(x)\beta_1$  and  $f_2(x)\beta_2$ . In particular we are interested in whether  $f_1(x)\beta_1$  is always greater than  $f_2(x)\beta_2$ . The test we will propose is a size  $\alpha$  test of

$$H_0: f_1(x)\beta_1 \leq f_2(x)\beta_2 \quad \text{for at least one } x \in R$$

vs.

$$H_A: f_1(x)\beta_1 > f_2(x)\beta_2 \quad \text{for every } x \in R.$$

Let  $b_1$  and  $b_2$  denote the least squares estimates of  $\beta_1$  and  $\beta_2$  and let

$$s^2 = \sum_{i=1}^2 \sum_{j=1}^{n_i} (y_{ij} - f_i(x_{ij})b_i)^2 / v$$

denote the pooled estimate of  $\sigma^2$  where  $v = n_1 - p_1 + n_2 - p_2$ .

The estimate  $b_i$  has a multivariate normal distribution with mean  $\beta_i$  and covariance matrix  $\sigma^2 D_i^{-1}$  where the  $(m, n)$  element of  $D_i$  is  $\sum_{j=1}^{n_i} f_{im}(x_{ij})f_{in}(x_{ij})$ . Let  $e(x) = f_1(x)D_1^{-1}f_1'(x) + f_2(x)D_2^{-1}f_2'(x)$ .

Then the variance of  $f_1(x)b_1 - f_2(x)b_2$  is  $\sigma^2 e(x)$ .

The test we will propose may be motivated in this way.

Consider comparing the two regression functions only at a single point  $x \in R$ . Let  $T_x = (f_1(x)b_1 - f_2(x)b_2)/\sqrt{e(x)}$ . The test which rejects  $H_{0x}$ :  $f_1(x)\beta_1 \leq f_2(x)\beta_2$  in favor of  $H_{Ax}$ :  $f_1(x)\beta_1 > f_2(x)\beta_2$  when  $T_x > t_{1-\alpha}(v)$  is a size  $\alpha$  test where  $t_{1-\alpha}(v)$  is the  $(1 - \alpha)$ th percentile of a t distribution with  $v$  degrees of freedom. The alternative of interest  $H_A$  is the intersection of all the  $H_{Ax}$  for all  $x \in R$ . It seems reasonable to decide in favor of  $H_A$  only if the test based on  $T_x$  decides in favor of  $H_{Ax}$  for every  $x \in R$ . This leads us to propose the following test.

Define the test statistic  $T$  by

$$(2.1) \quad T = \min_{x \in R} T_x.$$

Reject  $H_0$  in favor of  $H_A$  if and only if  $T > t_{1-\alpha}(v)$ . This test is always a level  $\alpha$  test in that the probability of a type one error is always less than or equal to  $\alpha$ . This test has size exactly equal to  $\alpha$  if the  $f_{ij}$  are continuous functions of  $x$  and there are values of  $\beta_1$  and  $\beta_2$  such that  $f_1(x)\beta_1 = f_2(x)\beta_2$  for one value of  $x \in R$  and  $f_1(x)\beta_1 > f_2(x)\beta_2$  for all other  $x \in R$ . These facts are proven in the Appendix. Section 3 contains examples of when the condition is satisfied and the size is exactly  $\alpha$ .

The test statistic  $T$  will usually have to be evaluated by numerical methods. This is discussed in Section 3. In some cases it can be evaluated explicitly. In particular it is shown in the Appendix that this test is equivalent to the tests proposed by Tsutakawa and Hewett (1978) and Hewett and Lababidi (1980) for the special models they consider if  $\alpha \leq .5$ .

## 2.2. A Numerical Application

We consider the data relating body and kidney weight for diabetic and healthy mice which was presented by Bishop (1973).

These data were analyzed by Tsutakawa and Hewett (1978). The data are plotted in Figure 1. The independent variable  $x$  is the body weight and the dependent variable  $y$  is the kidney weight. We consider modelling the mean of the kidney weight as a quadratic function of the body weight. We consider testing

$H_0: \beta_{11} + \beta_{12}x + \beta_{13}x^2 \leq \beta_{21} + \beta_{22}x + \beta_{23}x^2$  for some  $x \in R$   
versus  $H_A: \beta_{11} + \beta_{12}x + \beta_{13}x^2 > \beta_{21} + \beta_{22}x + \beta_{23}x^2$  for all  $x \in R$

where the first population is the diabetic population and

$R = \{x: 26 \leq x \leq 52\}$ . The least squares line for the 9 diabetic mice is given by  $1821.14 - 49.92x + .77x^2$ . The least squares line for the 25 healthy mice is given by

$-434.68 + 62.70x - .85x^2$ .  $s^2 = (66052 + 196505)/(9 - 3 + 25 - 3)$   
 $= 9377.0$ . The matrices  $D_1^{-1}$  and  $D_2^{-1}$  are

$$D_1^{-1} = \begin{bmatrix} 408.0894 & -19.1506 & .2214 \\ -19.1506 & .9031 & -.0105 \\ .2214 & -.0105 & .0001 \end{bmatrix}$$

and

$$D_2^{-1} = \begin{bmatrix} 81.1483 & -4.6954 & .0666 \\ -4.6954 & .2735 & -.0039 \\ .0666 & -.0039 & .0001 \end{bmatrix}$$

Thus  $T_x = a(x)/s\sqrt{e(x)}$  where  $a(x) = 2255.82 - 112.62x + 1.62x^2$   
and  $e(x) = 489.2377 - 47.6920x + 1.7526x^2 - .0288x^3 + .0002x^4$ .

Numerical minimization of  $T_x$  for  $x$  between 26 and 52 yields the test statistic  $T = .4797$  which corresponds to  $x = 40.10$ . This  $T$  is at approximately the 68th percentile of a  $t$  distribution with 28 degrees of freedom.

### 3. Properties of the Test

#### 3.1 Power Function Properties

The test we propose has size exactly  $\alpha$  if the  $f_{ij}$  are continuous functions and there are values of  $\beta_1$  and  $\beta_2$  such that  $f_1(x)\beta_1 = f_2(x)\beta_2$  for one value of  $x \in R$  and  $f_1(x)\beta_1 > f_2(x)\beta_2$  for all other  $x \in R$ . This condition is satisfied if the  $f_{ij}(x)$  include the constant 1, the linear functions  $x_i$ ,  $i = 1, \dots, k$ , and the quadratic functions  $x_i x_j$ ,  $i = 1, \dots, k$ ,  $j = 1, \dots, i$ .

Then  $\beta_1$  and  $\beta_2$  can be chosen so that

$f_1(x)\beta_1 - f_2(x)\beta_2 = (x - x_0)(x - x_0)'$ , the square of the distance from  $x$  to  $x_0$ , where  $x_0$  is a fixed element of  $R$ . For this choice,  $f_1(x)\beta_1 - f_2(x)\beta_2$  is zero for  $x = x_0$  and is positive for all other  $x$ . Another situation in which the condition is satisfied is if  $f_i(x)\beta_i = \beta_{i(k+1)} + \sum_{j=1}^k \beta_{ij}x_j$  and  $R = \{x: x_{j*} \leq x_j \leq x_j^*, j = 1, \dots, k\}$ , the model considered by Tsutakawa and Hewett (1978) and Hewett and Lababidi (1980).

The choice of  $\beta_2 = 0$ ,  $\beta_{11} = \dots = \beta_{1k} = 1$  and  $\beta_{1(k+1)} = -\sum_{j=1}^k x_{j*}$  yields  $f_1(x)\beta_1 - f_2(x)\beta_2 = \sum_{j=1}^k (x_j - x_{j*})$  which is zero for  $x = (x_{1*}, \dots, x_{k*})$  and positive for all other  $x \in R$ . A numerical corroboration of the fact that the size is exactly  $\alpha$  is found in the following simulation study.

A simulation study was conducted to investigate the power function of the test. In this study the regression functions were  $f_i(x)\beta_i = \beta_{i1} + \beta_{i2}x + \beta_{i3}x^2$ ,  $i = 1, 2$ . The variance  $\sigma^2$  was set equal to one. The sample sizes  $n_1$  and  $n_2$  were both 10 with 3 observations at each of  $x = 1$  and  $x = -1$  and 4 observations at  $x = 0$ .  $R = \{x: -1 \leq x \leq 1\}$ . The size of the test was fixed at  $\alpha = .05$  by using  $t_{.95}(14) = 1.761$ . The International Mathematics and Statistics Library programs GGNSM and GGCHS were used to generate the random vector  $b_1 - b_2$  and the random variable  $S^2$ . A total of 3000 repetitions were used to obtain

each of the estimates in Tables 1 and 2.

The maximum probability of a Type I error takes place when  $f_1(x)\beta_1 = f_2(x)\beta_2$  for one  $x$  and  $f_1(x)\beta_1 - f_2(x)\beta_2$  becomes large for all other  $x$ . This can be observed in Table 1 where the probability of a Type I error is given for various values of  $\beta_1$  and  $\beta_2$  in the null hypothesis. As one proceeds down Columns II, III, IV or V of Table 1,  $f_1(x)\beta_1 = f_2(x)\beta_2$  for one value of  $x$  ( $x = -1$  for Columns III, IV and V and  $x = 0$  for Column II) and  $f_1(x)\beta_1 - f_2(x)\beta_2$  is becoming large for all other values of  $x$ . The probability of a Type I error increases to  $\alpha = .05$  as one proceeds down any column. The estimates slightly exceed .05 in a few cases due to sampling error.

The power function of the proposed test exhibits the following monotonicity property. If  $(\beta_1, \beta_2)$  and  $(\beta_1^*, \beta_2^*)$  are two parameter vectors which satisfy

$$(3.1) \quad f_1(x)\beta_1^* - f_2(x)\beta_2^* \geq f_1(x)\beta_1 - f_2(x)\beta_2$$

for every  $x$  with strict inequality for some  $x$ , then the power at  $(\beta_1^*, \beta_2^*)$  is greater than the power at  $(\beta_1, \beta_2)$ . This property is apparent as one proceeds down any column in Tables 1 or 2, across the first three columns in any row of Table 2, or across the last three columns in any row of Table 1. It is also apparent if one compares any two corresponding entries in Table 2a and 2b since the regression functions are farther apart in 2b than in 2a.

The power of the test is near one only if  $f_1(x)\beta_1 - f_2(x)\beta_2$  is large for all  $x$ . This is the case for the lower entries in Table 2. In Table 2, the minimum distance between the regression functions is  $c$ . The power nears one only as the minimum distance  $c$  becomes large.

The test we propose is biased in that the probability of rejecting  $H_0$  is less than  $\alpha$  for some  $(\beta_1, \beta_2)$  in  $H_A$ . The feature was noted by Tsutakawa and Hewett (1978) for the special model they considered and it continues to exist for the more general models we consider. This biasedness can be observed in the entries for  $c = .5$  which are less than .95 in Table 2. But as noted by Tsutakawa and Hewett (1978) for their special case, the test we propose is consistent in that, for any fixed point  $(\beta_1, \beta_2)$  in  $H_A$ , the power can be made arbitrarily near one by choosing the sample sizes sufficiently large. Although we do not feel this bias is serious, it should be noted that the power of the test we propose may be small if  $f_1(x)\beta_1$  exceeds  $f_2(x)\beta_2$  by only a small amount over most of  $R$ .

The power function properties we have described in this section are true in general, not just for the case of quadratic regression we considered in the simulation experiment. The proofs of these facts can be accomplished using the methods employed in proofs in the Appendix.

### 3.2. Computational Shortcuts

In (2.1) the test statistic  $T$  was defined as the minimum of  $T_x$  over the set  $R$ . Typically the computation of the test statistic will be accomplished by a numerical minimization of  $T_x$ . But to perform the test the actual value of  $T$  need not be computed. One only needs to know whether  $T > t_{1-\alpha}(v)$  or  $T \leq t_{1-\alpha}(v)$ . In this section we describe two shortcuts which allow the determination of whether  $T > t_{1-\alpha}(v)$  or  $T \leq t_{1-\alpha}(v)$  without the actual computation of  $T$ .

#### 3.2.1. Shortcut for determining if $H_0$ is accepted.

Let  $X^*$  denote an arbitrary finite subset of  $R$ . For example, if  $R = \{x: x_{i*} \leq x_i \leq x_i^*, i = 1, \dots, k\}$ ,  $X^*$  might be the set of  $2^k$  extreme points  $(x_1, \dots, x_k)$  where  $x_i = x_{i*}$  or  $x_i = x_i^*$ . Let  $T' = \min_{x \in X^*} T_x$ . Since  $T \leq T'$ , if  $T' \leq t_{1-\alpha}(v)$  accept  $H_0$ . Furthermore if  $T' = t'$  the significance probability associated with  $T$  is at least  $P(T_0 > t')$  where  $T_0$  has a central t distribution with  $v$  degrees of freedom.

For the body kidney weight data of Section 2.2,

$R = \{x: 26 \leq x \leq 52\}$ .  $T_x$  for  $x = 26$  is .9962 and  $T_x$  for  $x = 52$  is .6347. Either one of these points is less than  $T_{.95}(28) = 1.7011$  so the test accepts  $H_0$  at level .05. Furthermore the significance probability associated with  $T$  is at least  $P(T_0 \geq .6347) = .27$ .

The exact significance probability computed in Section 2.2 was .32.

In Tables 1, and 2 the second (middle) number for each entry is the proportion of the acceptances which were determined by the shortcut method. These values indicate that the usefulness of this shortcut depends on the actual regression functions. But, in many cases, a large proportion of the acceptances were determined by this shortcut. In 18 out of the 62 cases in which there were some acceptances, all of the acceptances were determined by the shortcut.

### 3.2.2. Shortcut for determining if $H_0$ is rejected.

For this shortcut to be valid,  $\alpha$  must be no more than .5.

Since  $\alpha$  usually satisfies  $\alpha \leq .1$ , this restriction is not practically important. Let  $m$  denote the number of distinct non-constant functions in  $\{f_{ij}(x): i = 1, 2; j = 1, \dots, p_i\}$ . Let  $Z^*$  denote the set of  $2^m$  points

$(z_1^*, z_2^*) = ((z_{11}, \dots, z_{1p_1}), (z_{21}, \dots, z_{2p_2}))$  formed by replacing  $f_{ij}(x)$  by either  $\max_{x \in R} f_{ij}(x)$  or  $\min_{x \in R} f_{ij}(x)$  in  $(f_1(x), f_2(x))$ . Note that if  $f_{1r}(x) = f_{2s}(x)$  then  $z_{1r} = z_{2s}$ , i.e., the maximum or minimum is used on both  $z_1^*$  and  $z_2^*$ . Let

$$T^* = \min_{z^* \in Z^*} T_{z^*} \text{ where}$$

$T_{z^*} = (z_{11}^* b_1 - z_{21}^* b_2) / s \sqrt{z_{11}^{*2} D_1^{-1} z_{11}^{**} + z_{22}^{*2} D_2^{-1} z_{22}^{**}}$ . If  $T^* > t_{1-\alpha}(v)$  then  $T > t_{1-\alpha}(v)$  and  $H_0$  can be rejected. Furthermore, if

$T^* = t^*$ , the significance probability associated with  $T$  is less than or equal to  $P(T_0 > t^*)$  where  $T_0$  has a central  $T$  distribution with  $v$  degrees of freedom. If  $T^* = T_{z^*}$  where  $z^* = (f_1(x), f_2(x))$

for some  $x \in R$  then, in fact, the significance probability equals  $P(T_0 > t^*)$ .

For example, suppose  $f_1(x) = (1, x, x^2)$ , and  $f_2(x) = (1, x)$ , and  $R = \{x: -1 \leq x \leq 2\}$ . There are two distinct nonconstant functions,  $x$  and  $x^2$ , and the  $2^2 = 4$  values of  $(z_1^*, z_2^*)$  in  $Z^*$  are  $((1, -1, 0), (1, -1)), ((1, -1, 4), (1, -1)), ((1, 2, 0), (1, 2))$  and  $((1, 2, 4), (1, 2))$ . Points like  $((1, -1, 0), (1, 2))$  where  $x$  has been replaced by its minimum in  $z_1^*$  and its maximum in  $z_2^*$  are not in  $Z^*$ .

The validity of this shortcut is based on two facts, the fact about functions which are the ratios of linear functions and square roots of positive quadratic functions mentioned in the proof of Theorem 3 and the fact that  $A = \{(f_1(x), f_2(x)): x \in R\}$  is a subset of

$$B = \{(z_1, z_2): \min_{x \in R} f_{ij}(x) \leq z_{ij} \leq \max_{x \in R} f_{ij}(x) \text{ and } z_{1r} = z_{2s} \text{ if } f_{1r}(x) = f_{2s}(x)\}$$

so a minimum over  $A$  is not less than a minimum over  $B$ .

This shortcut was used in the simulation study of Section 3.1. In this case  $R = \{x: -1 \leq x \leq 1\}$ .  $f_1(x) = f_2(x) = (1, x, x^2)$  so there are  $m = 2$  distinct nonconstant functions. The  $2^m = 4$  points in  $Z^*$  are  $((1, -1, 0), (1, -1, 0)), ((1, -1, 1), (1, -1, 1)), ((1, 1, 0), (1, 1, 0))$  and  $((1, 1, 1), (1, 1, 1))$ .

In Tables 1 and 2, the third (bottom) number for each entry is the proportion of the rejections which were detected by this

shortcut. The usefulness of this shortcut is seen to depend on the actual value of the regression function but in many cases the proportion is fairly high. In 17 of 71 cases all of the rejections were detected by this method, avoiding numerical minimization of  $T_x$ .

### 3.2.3. Numerical minimization of $T_x$ .

If neither of the shortcuts presented in Sections 3.2.1 or 3.2.2 determine whether  $H_0$  is to be accepted or rejected or if the exact value of  $T$  is desired to compute an exact significance probability, then the function  $T_x$  must be minimized by numerical methods to determine the value of the test statistic  $T$ . The problem of minimizing a function such as  $T_x$  which is the ratio of two functions of  $x$  has been studied extensively in the mathematical programming literature by Charnes and Cooper (1962), Swarup (1965) Sharma (1967) and Craven and Mond (1973, 1975a, and 1975b). These authors have found that this non-linear programming problem is equivalent to other nonlinear programming problems which do not involve fractions. These results could simplify the numerical minimization of  $T_x$ .

## 4. Comparison of a Regression Function with a Specified Function.

The hypothesis that the regression function from a population is everywhere above a specified function can be tested with a test similar to the one described in Section 2. For this problem we only have the sample  $\{(X_{1j}, Y_{1j}); j = 1, \dots, n_1\}$  from

the first population. We wish to test

$$H_0: f(x)\beta_1 \leq g(x) \quad \text{for at least one } x \in R$$

vs

$$H_A: f(x)\beta_1 > g(x) \quad \text{for all } x \in R$$

where  $g(x)$  is a specified function. The function  $g(x)$  may be a constant  $c$  in which case we are testing whether the mean of the response is greater than  $c$  for all possible values of the independent variable  $x$ . Let

$$s_1^2 = \sum_{j=1}^{n_1} (y_{1j} - f_1(x_{1j})\beta_1)^2 / v_1$$

where  $v_1 = n_1 - p_1$ . Let  $e_1(x) = f_1(x)D_1^{-1}f_1'(x)$  and  $T_{1x} = (f_1(x)\beta_1 - g(x))/s_1 \sqrt{e_1(x)}$ . Define the test statistic  $T_1$  by

$$T_1(x) = \min_{x \in R} T_{1x}.$$

A level  $\alpha$  test of  $H_0$  verses  $H_A$  is given by reject  $H_0$  if  $T_1 > t_{1-\alpha}(v_1)$ .

This test enjoys all the same properties as the test based on  $T$  described in Sections 2 and 3. For example, the test has size exactly  $\alpha$  if  $f_{11}, \dots, f_{1p_1}$  and  $g$  are all continuous functions and there is a value of  $\beta_1$  such that  $f(x)\beta_1 = g(x)$  for one value of  $x \in R$  and  $f_1(x)\beta_1 > g(x)$  for all other  $x \in R$ . The proofs of these properties are analogous to those for  $T$  with the function  $f_1(x)\beta_1 - f_2(x)\beta_2$  replaced by  $f_1(x)\beta_1 - g(x)$ . These proofs are not given herein.

Table 1

Power of the Test and Percentage of Acceptances and Rejections  
by Shortcuts<sup>1</sup> for Selected Points in  $H_0^2$ .

c	I	II	III	IV	V
0	.0003	.0003	.0003	.0003	.0003
	99	99	99	99	99
	100	100	100	100	100
1	.0027	.0080	.0037	.0067	.0087
	100	90	98	98	99
	50	33	27	50	50
2	.0053	.0287	.0110	.0237	.0327
	100	40	97	98	99
	75	22	30	39	63
5	.0060	.0483	.0267	.0493	.0523
	100	0	97	100	100
	100	16	16	32	78
25	.0060	.0510	.0460	.0523	.0523
	100	0	99	100	100
	100	15	9	30	100
1000	.0060	.0510	.0523	.0523	.0523
	100	0	100	100	100
	100	15	6	30	100

<sup>1</sup>First (top) entry: estimated power of the test  
Second (middle) entry: percentage of acceptances detected by  
shortcut in Section 3.2.1  
Third (bottom) entry : percentage of rejections detected by  
shortcut in Section 3.2.2

$$\begin{aligned}
 ^2 \text{Column I : } & f_1(x)\beta_1 - f_2(x)\beta_2 = c(1 - x^2) \\
 \text{Column II : } & f_1(x)\beta_1 - f_2(x)\beta_2 = cx^2 \\
 \text{Column III: } & f_1(x)\beta_1 - f_2(x)\beta_2 = c(x + 1)^2/4 \\
 \text{Column IV : } & f_1(x)\beta_1 - f_2(x)\beta_2 = c(x + 1)/2 \\
 \text{Column V : } & f_1(x)\beta_1 - f_2(x)\beta_2 = c(-x^2 + 2x + 3)/4
 \end{aligned}$$

Table 2a

Power of the Test and Percentage of Acceptances and Rejections  
by Shortcuts<sup>1</sup> for Selected Points in  $H_A^2$

c	I	II	III	IV	V
.5	.0053	.0290	.0547	.0180	.0560
	98	94	96	99	73
	38	33	51	76	21
1	.0510	.1460	.2023	.0990	.2327
	94	87	94	99	51
	45	44	63	78	36
2	.5197	.6897	.7357	.6030	.7967
	82	81	96	99	30
	65	65	81	92	59
3	.9323	.9647	.9667	.9387	.9907
	91	93	99	100	29
	90	90	97	100	88
5	1.000	1.000	1.000	1.000	1.000
	--	--	--	--	--
	100	100	100	100	100

<sup>1</sup>See Table 1 footnote

<sup>2</sup>Column I :  $f_1(x)\beta_1 - f_2(x)\beta_2 = c$

Column II :  $f_1(x)\beta_1 - f_2(x)\beta_2 = (x + 1)^2/4 + c$

Column III :  $f_1(x)\beta_1 - f_2(x)\beta_2 = (-x^2 + 2x + 3)/4 + c$

Column IV :  $f_1(x)\beta_1 - f_2(x)\beta_2 = -x^2 + 1 + c$

Column V :  $f_1(x)\beta_1 - f_2(x)\beta_2 = x^2 + c$

Table 2b

Power of the Test and Percentage of Acceptances and Rejections  
by Shortcuts<sup>1</sup> for Selected Points in  $H_A^2$ .

c	I	II	III	IV	V
.5	.0053	.0870	.1453	.0250	.1510
	98	94	100	100	0
	38	11	79	100	17
1	.0510	.2543	.3167	.1107	.3680
	94	92	100	100	0
	45	15	87	100	28
2	.5197	.7557	.7700	.6077	.8460
	82	94	100	100	0
	65	32	96	100	57
3	.9323	.9677	.9680	.9387	.9933
	91	99	100	100	0
	90	68	100	100	88
5	1.0000	1.0000	1.0000	1.0000	1.0000
	--	--	--	--	--
	100	100	100	100	100

<sup>1</sup>See Table 1 footnote.

<sup>2</sup>Column I :  $f_1(x)\beta_1 - f_2(x)\beta_2 = c$   
 Column II :  $f_1(x)\beta_1 - f_2(x)\beta_2 = (x + 1)^2 + c$   
 Column III:  $f_1(x)\beta_1 - f_2(x)\beta_2 = -x^2 + 2x + 3 + c$   
 Column IV :  $f_1(x)\beta_1 - f_2(x)\beta_2 = -4x^2 + 4 + c$   
 Column V :  $f_1(x)\beta_1 - f_2(x)\beta_2 = 4x^2 + c$

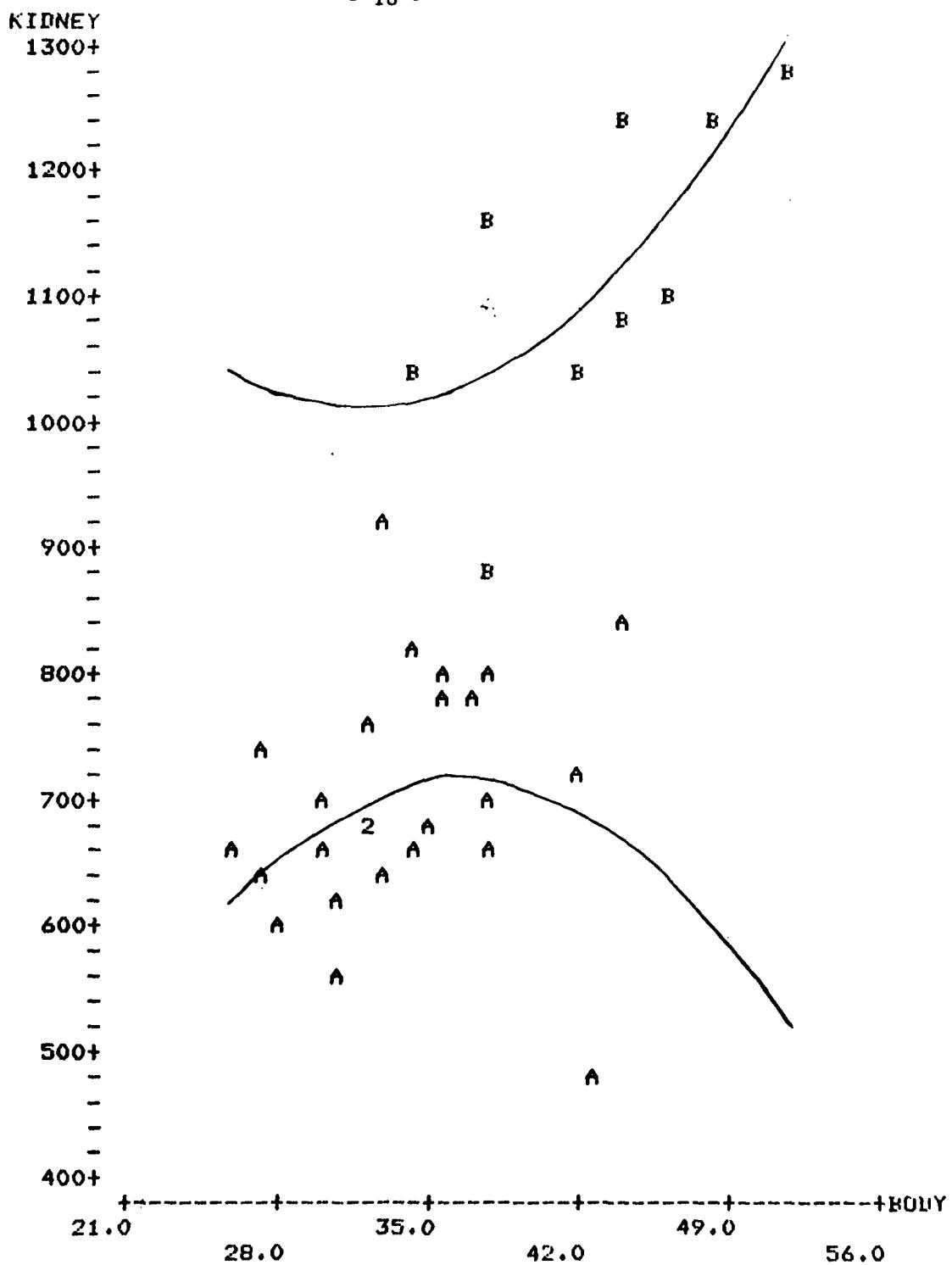


Figure 1  
Body and Kidney Weight for  
Healthy (A) and Diabetic (B) Mice

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Appendix

Proofs regarding the size of the test and the equivalence of the test to the test proposed by Tsutakawa and Hewett (1978) and Hewett and Lababidi (1980) are given in this appendix.

The size of the test.

Theorem 1: Under the assumptions of our model the test has level  $\alpha$ , i.e.,

$$A.1 \quad \sup_{(\beta_1, \beta_2) \in H_0} P_{\beta_1, \beta_2} (T > t_{1-\alpha}(v)) \leq \alpha.$$

Proof: Fix  $(\beta_1, \beta_2) \in H_0$ . There is an  $x_0 \in R$  such that  $f_1(x_0)\beta_1 \leq f_2(x_0)\beta_2$ . Then, with probability one,  $T \leq T_{x_0} \leq Q$  where

$$Q = \frac{f_1(x_0)\beta_1 - f_2(x_0)\beta_2 - (f_1(x_0)\beta_1 - f_2(x_0)\beta_2)}{S\sqrt{e(x_0)}}$$

$Q$  has a  $t$  distribution with  $v$  degrees of freedom. So

$$P_{\beta_1, \beta_2} (T > t_{1-\alpha}(v)) \leq P(Q > t_{1-\alpha}(v)) = \alpha.$$

Since  $\beta_1$  and  $\beta_2$  were arbitrary, A.1 is true. ||

Theorem 2: Suppose that all the  $f_{ij}(x)$ ,  $i = 1, 2$ ,  $j = 1, \dots, p_i$ , are continuous on  $R$ . If there exist  $\beta_1^*$  and  $\beta_2^*$  such that  $f_1(x_0)\beta_1^* = f_2(x_0)\beta_2^*$  for one  $x_0 \in R$  and  $f_1(x)\beta_1^* > f_2(x)\beta_2^*$  for all other  $x \in R$  then the test has size exactly  $\alpha$ , i.e.,

$$\sup_{(\beta_1, \beta_2) \in H_0} P_{\beta_1, \beta_2} (T > t_{1-\alpha}(v)) = \alpha.$$

The proof of Theorem 2 will use Lemma 1 which can be proved using standard analysis methods.

Lemma 1: Let  $g_n(x)$ ,  $n = 1, 2, \dots$ , be continuous functions on a compact set  $R$ . Suppose there exists an  $x_0 \in R$  such that  $g_n(x_0)$  is constant (say  $c$ ) for all  $n$ . Suppose  $g_n(x)$  increases to infinity as  $n \rightarrow \infty$  for all  $x \neq x_0$ . Then

$$(A.2) \quad \lim_{n \rightarrow \infty} \min_{x \in R} g_n(x) = c.$$

Proof of Theorem 2: By Theorem 1 it suffices to show there exists a sequence  $(\beta_1^n, \beta_2^n)$ ,  $n = 1, 2, \dots$ , such that,  $(\beta_1^n, \beta_2^n) \in H_0$  for  $n = 1, 2, \dots$ , and

$$(A.4) \quad \lim_{n \rightarrow \infty} P_{\beta_1^n, \beta_2^n} (T > t_{1-\alpha}(v)) \geq \alpha.$$

The estimates  $b_i$ ,  $i = 1, 2$ , can be written as  $b_i = z_i + \beta_i$  where  $z_1$ ,  $z_2$  and  $S$  are independent, and  $z_i$  has an  $p_i$ -variate normal distribution with mean 0 and variance-covariance matrix  $\sigma_{D_i}^{2-1}$ . In terms of these quantities, the statistics  $T$  and  $T_x$  can be written as

$$T = T(z_1, z_2, S, \beta_1, \beta_2) = \min_{x \in R} T_x(z_1, z_2, S, \beta_1, \beta_2)$$

and

$$T_x(z_1, z_2, S, \beta_1, \beta_2) = \frac{f_1(x)z_1 - f_2(x)z_2 + f_1(x)\beta_1 - f_2(x)\beta_2}{S^e(x)}.$$

Consider the sequence  $(\beta_1^n, \beta_2^n)$  defined by  $\beta_i^n = n\beta_i^*$  where the  $\beta_i^*$  are defined in the statement of Theorem 2. For a fixed value of  $z_1 \in R^{p_1}$ ,  $z_2 \in R^{p_2}$  and  $s > 0$ , define

$$g_n(x) = T_x(z_1, z_2, s, \beta_1^n, \beta_2^n).$$

The  $g_n(x)$  satisfy the conditions of Lemma 1 since 1)  $f_{ij}$  are continuous, 2)  $s\sqrt{e(x)} > 0$ , 3)  $f_1(x_0)\beta_1^n = f_2(x_0)\beta_2^n$  and 4)  $f_1(x)\beta_1^n - f_2(x)\beta_2^n$  increases to infinity as  $n \rightarrow \infty$  for all  $x \neq x_0$ . By Lemma 2,  $\lim_{n \rightarrow \infty} T(z_1, z_2, s, \beta_1^n, \beta_2^n) = T_{x_0}(z_1, z_2, s, \beta_1^*, \beta_2^*)$ . Since  $z_1$ ,  $z_2$ , and  $s$  were arbitrary, this implies that  $T(z_1, z_2, s, \beta_1^n, \beta_2^n)$  converges to  $T_{x_0}(z_1, z_2, s, \beta_1^*, \beta_2^*)$  with probability one and hence in distribution. Thus

$$\begin{aligned} \lim_{n \rightarrow \infty} P_{\beta_1^n, \beta_2^n}(T > t_{1-\alpha}(v)) &= \lim_{n \rightarrow \infty} P(T(z_1, z_2, s, \beta_1^n, \beta_2^n) > t_{1-\alpha}(v)) \\ &= P(T_{x_0}(z_1, z_2, s, \beta_1^*, \beta_2^*) > t_{1-\alpha}(v)) \\ &= \alpha. \quad || \end{aligned}$$

#### Equivalence with tests proposed by Tsutakawa and Hewett (1978)

and Hewett and Latabidi (1980).

Theorem 3: Suppose  $f_i(x)\beta_i = \beta_{i0} + \sum_{j=1}^k \beta_{ij} x_j$  and  $R$  has the form  $R = \{x: x_{j*} \leq x_j \leq x_j^*, j = 1, \dots, k\}$ . Consider the test which rejects  $H_0$  if  $T^* > t_{1-\alpha}(v)$  where  $T^* = \min_{x \in X^*} T_x$  and  $X^*$  is the set of  $2^k$  points for which  $x_j$  is either  $x_{j*}$  or  $x_j^*$ . Suppose  $\alpha \leq .5$ . Then the tests based on  $T^*$  and  $T$  are equivalent.

Proof. For any  $k + 1$  dimensional vectors  $b_1$  and  $b_2$  and  $s > 0$ ,  $T_x$  is a linear function of  $(x_1, \dots, x_k)$  divided by the square root of a quadratic function of  $(x_1, \dots, x_k)$  which is positive for all  $(x_1, \dots, x_k) \in \mathbb{R}^k$ . Such a function has the property that  $T^* = \min_{x \in X^*} T_x \geq 0$  implies  $T^* = \min_{x \in R} T_x = T$ . (This is easily proved for  $k = 1$  and can be proven for general  $k$  by induction.). For any  $b_1, b_2$  and  $s > 0$ ,  $T \leq T^*$  so if  $T$  rejects  $H_0$ , so does  $T^*$ . Suppose  $b_1, b_2$  and  $s$  are such that  $T^*$  rejects  $H_0$ . Then  $T^* > t_{1-\alpha}(v) \geq 0$ , since  $\alpha \leq .5$ , so  $T = T^*$  and  $T$  also rejects  $H_0$ . ||

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SECURITY CLASSIFICATION OF THIS PAGE

REPORT DOCUMENTATION PAGE

1. REPORT NUMBER FSU Statistics M607 USARO D-52	2. GOVT ACCESSION NO.	3. RECIPIENT'S CATALOG NUMBER
4. TITLE (and subtitle) Testing Whether One Regression Function is Everywhere Larger Than Another	5. TYPE OF REPORT & PERIOD COVERED Technical	
6. PERFORMING ORGANIZATION NAME AND ADDRESS Department of Statistics Florida State University Tallahassee, FL 32306	7. AUTHOR(S) Roger L. Berger	
8. CONTRACT OR GRANT NUMBER(s) USARO Grant DAAG29-79-C-0158	9. PROGRAM ELEMENT, PROJECT, TASK AREA & WORK UNIT NUMBERS	
10. CONTROLLING OFFICE NAME AND ADDRESS U.S. Army Research Office P.O. Box 12211 Research Triangle Park, N.C. 27709	11. REPORT DATE February, 1982	12. NUMBER OF PAGES 23
13. MONITORING AGENCY NAME & ADDRESS (if different from Controlling Office)	14. SECURITY CLASS. (of this report) Unclassified	
15. DECLASSIFICATION/DEGRADED SCHEDULE		
16. DISTRIBUTION STATEMENT (of this report) Distribution unlimited		
17. DISTRIBUTION STATEMENT (of the abstract entered in Block 20, if different from report		
18. SUPPLEMENTARY NOTES <small>(THE VIEW, OPINIONS, AND/OR FINDINGS CONTAINED IN THIS REPORT ARE THOSE OF THE AUTHOR(S) AND SHOULD NOT BE CONSTRUED AS AN OFFICIAL DEPARTMENT OF THE ARMY POSITION, POLICY, OR DE- CISION, UNLESS SO DESIGNATED BY OTHER DOCUMENTATION.)</small>		
19. KEY WORDS		
20. ABSTRACT (Continue on reverse side if necessary and identify by block number) <p>The problem of testing whether one regression function is larger than another on a specified compact set <math>R</math> is considered. The regression functions must be linear functions of the parameters but need not be linear functions of the independent variables. The proposed test statistic is compared to a standard <math>t</math> percentile. The test has an exactly specified size in typical situations. Properties of the power function of the test are investigated. The related question of comparing a regression function to a specified function is also considered.</p>		

